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L4
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L5
            689 S L2 AND L4
                SAV L5 HOPE753/A
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            102 S E3, E13
L8
                E DAVIDSON DON/AU
             62 S E3, E5, E6, E12, E13
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                E WANG J/AU
            270 S E55-E60
L10
                E WANG JI/AU
L11
             94 S E3, E18
                E WANG JIE/AU
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            648 S E3
                E WANG JIEYI/AU
L13
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                E GUBBINS E/AU
L14
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L15
           9014 S E3, E4 OR ABBOT?/PA, CS
L16
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L17
              7 S L8-L15 AND L6
L18
              7 S L16, L17
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                E E4+ALL
          14221 S E5+NT
L23
           1247 S E24, E32, E33, E42, E43, E49, E50
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                E E63+ALL
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              8 S L32 AND PLASMINOGEN
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             13 S L32, L34, L35
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L38
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SAV L38 HOPE753A/A

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FILE COVERS 1907 - 15 Feb 2005 VOL 142 ISS 8 FILE LAST UPDATED: 14 Feb 2005 (20050214/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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- L36 ANSWER 1 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
- 2004:176555 HCAPLUS
- 140:229440 DN
- Entered STN: 04 Mar 2004 .ED
- Antiangiogenic peptides derived from mammalian protein kringle 5 and use for treating angiogenic diseases
- Davidson, Donald J.
- Abbott Laboratories, USA PΑ
- U.S., 47 pp., Cont.-in-part of U.S. Ser. No. 851,350. CODEN: USXXAM
- DTPatent
- English LA
- IC ICM A61K038-00
- 514012000; 514002000; 514013000; 514014000; 514015000; 514016000; 514017000; 514018000; 514648000; 514336000
- 1-8 (Pharmacology)

Section cross-reference(s): 3, 6, 13

FAN. CNT 4							
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US 20	04138127	A1	20040715	US	2004-753646	20040108	<
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US 19	97-851350	A2	19970505				
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CLASS							

US 6699838	ICM	A61K038-00				
	NCL	514012000;	514002000;	514013000;	514014000;	514015000;
		514016000;	514017000;	514018000;	514648000;	514336000

CLASS PATENT FAMILY CLASSIFICATION CODES

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US 5981484
                ECLA
                        C12N009/68
US 6057122
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                ECLA
                        C12N009/68
US 2004138127
                                                                             <--
    The invention provides mammalian kringle 5 fragments and kringle 5 fusion
AB
    proteins as a compds. for treating angiogenic diseases. The
     invention also provides methods and compns. for inhibiting
     angiogenic diseases.
ST
     antiangiogenic peptide human protein kringle
     angiogenesis
IT
     Angiogenesis inhibitors
     Human
     Protein sequences
        (antiangiogenic peptides derived from mammalian protein
        kringle 5 and use for treating angiogenic diseases)
IT
     Peptides, biological studies
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (antiangiogenic peptides derived from mammalian protein
        kringle 5 and use for treating angiogenic diseases)
IT
     Fusion proteins (chimeric proteins)
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (k5; antiangiogenic peptides derived from mammalian protein
        kringle 5 and use for treating angiogenic diseases)
IT
     Protein motifs
        (kringles; antiangiogenic peptides derived from mammalian
        protein kringle 5 and use for treating angiogenic diseases)
     666829-12-5P 666867-41-0P
                                666867-42-1P
                                                666867-43-2P
IT
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (antiangiogenic peptide sequence; antiangiogenic
        peptides derived from mammalian protein kringle 5 and use for treating
        angiogenic diseases)
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                                 666870-08-2
                                               666870-09-3
                                                              666870-10-6
TT
     666870-06-0
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                                               666870-34-4
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        (unclaimed nucleotide sequence; antiangiogenic peptides
        derived from mammalian protein kringle 5 and use for treating
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     666870-37-7 666870-38-8 666870-39-9
     RL: PRP (Properties)
        (unclaimed protein sequence; antiangiogenic peptides derived
        from mammalian protein kringle 5 and use for treating
        angiogenic diseases)
RE.CNT
              THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD
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MEDLINE

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- L36 ANSWER 2 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
- AN 2004:99288 HCAPLUS
- DN 140:331802
- ED Entered STN: 06 Feb 2004
- TI Lysyl 4-aminobenzoic acid derivatives as potent small molecule mimetics of plasminogen kringle 5
- AU Sheppard, George S.; Kawai, Megumi; Craig, Richard A.; Davidson, Donald J.; Majest, Sandra M.; Bell, Randy L.; Henkin, Jack
- CS Abbott Laboratories, Global Pharmaceutical Research and Development, Abbott Park, IL, 60064, USA
- SO Bioorganic & Medicinal Chemistry Letters (2004), 14(4), 965-966 CODEN: BMCLE8; ISSN: 0960-894X
- PB Elsevier Science B.V.
- DT Journal
- LA English
- CC 1-3 (Pharmacology)
 Section cross-reference(s): 34
- OS CASREACT 140:331802
- AB Kringle 5, a proteolytic fragment of human plasminogen has been shown to potently inhibit angiogenesis. The tetrapeptide KLYD derived from kringle 5 has been shown to capture many activities of kringle 5 in vitro. Further simplification has been achieved by replacement of the two central amino acids with a 4-aminobenzoic acid spacer group. Mols. displaying the required recognition groups on this core show similar in vitro properties to kringle 5, and are able to displace radiolabeled protein from a high affinity binding site on endothelial cells.
- ST lysyl aminobenzoic acid deriv prepn plasminogen kringle mimetic angiogenesis
- IT Blood vessel
 - (endothelium; preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of **plasminogen** kringle 5 that inhibit HMVEC chemotaxis)
- IT Protein motifs
 - (kringles; preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of **plasminogen** kringle 5 that inhibit HMVEC chemotaxis)
- IT Angiogenesis inhibitors

Chemotaxis

Human

Peptidomimetics

Structure-activity relationship (preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of plasminogen kringle 5 that inhibit HMVEC chemotaxis) IT Endothelium (vascular; preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of plasminogen kringle 5 that inhibit HMVEC chemotaxis) 9001-91-6, Plasminogen IT RL: BSU (Biological study, unclassified); BIOL (Biological study) (preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of plasminogen kringle 5 that inhibit HMVEC chemotaxis) 250789-27-6P 250789-59-4P 250789-78-7P TT 250789-79-8P 679784-35-1P 679784-36-2P 679784-37-3P 679784-38-4P RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of plasminogen kringle 5 that inhibit HMVEC chemotaxis) TT679784-39-5 RL: PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of plasminogen kringle 5 that inhibit HMVEC chemotaxis) IT 692-04-6 1155-64-2 2389-45-9 4530-20-5 6404-28-0 13795-73-8, L-Aspartic acid, bis(1,1-dimethylethyl) ester 18144-47-3, tert-Butyl 4-aminobenzoate 21887-64-9 156682-54-1, 3-(Benzyloxy)phenylboronic 250790-07-9 RL: RCT (Reactant); RACT (Reactant or reagent) (preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of plasminogen kringle 5 that inhibit HMVEC chemotaxis) IT250790-08-0P 679784-40-8P 679784-41-9P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of lysyl 4-aminobenzoic acid derivs. as potent small mol. mimetics of plasminogen kringle 5 that inhibit HMVEC chemotaxis) THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT RE (1) Cao, Y; J Biol Chem 1997, V272, P22924 HCAPLUS (2) Davidson, D; 215th ACS National Meeting 1998, Abstract MEDI-207 (3) Davidson, D; Proc Am Assoc Cancer Res 2000, V41, P486 (4) Hanahan, D; Cell 1996, V86, P353 HCAPLUS (5) Majest, S; Proc Am Assoc Cancer Res 2001, V42, P484 (6) Pozdnev, V; Int J Pept Prot Res 1994, V44, P36 HCAPLUS (7) Qian, Y; J Biol Chem 1994, V269, P12410 HCAPLUS L36 ANSWER 3 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN 2003:448045 HCAPLUS AN139:30780 DN ED Entered STN: 11 Jun 2003 Methods and compositions for generating angiostatin TI Soff, Gerald; Gately, Stephen T.; Twardowski, Przemyslaw IN PANorthwestern University, USA U.S., 46 pp., Cont.-in-part of U.S. Ser. No. 710,305. SO CODEN: USXXAM DT Patent

LA

IC

English

ICM A61K038-00

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NCL 514012000; 435217000; 530350000; 530380000
     1-6 (Pharmacology)
    Section cross-reference(s): 2
FAN.CNT 2
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    US 6576609
                       B1 20030610 US 1997-991761 19971216 <--
PΙ
                       A 19980901 US 1996-710305 19960917
A1 19980416 WO 1997-US16539 19970917 <--
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    US 5801012
    WO 9815574
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            KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG,
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            GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
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PRAI US 1996-710305
     WO 1997-US16539
                        A1
                              19970917
CLASS
               CLASS PATENT FAMILY CLASSIFICATION CODES
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 US 6576609
              ICM A61K038-00
                       514012000; 435217000; 530350000; 530380000
               NCL
              ECLA
                      A61K038/16B1+M; A61K038/49+M; C12N009/68
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                       A61K038/16B1+M; A61K038/49+M; C12N009/68
 US 5801012
               ECLA A61K038/16B1+M; A61K038/49+M; C12N009/68
 WO 9815574
     The invention provides a method of treating a neoplastic disease in a
AB
     human by administering a therapeutically effective amount of
     plasminogen activator effective to increase the amount of
     angiostatin present in the human to treat the disease. The invention also
     provides a method of treating a neoplastic disease in a human by
     administering a therapeutically effective amount of plasminogen
     activator and sulfhydryl donor effective to increase the amount of
     angiostatin present in the human to treat said disease.
     angiostatin generation plasminogen activator sulfhydryl donor;
ST
     neoplasm treatment angiostatin generation plasminogen activator
ΙT
     Sulfhydryl group
        (donors; generating angiostatin using plasminogen activator
        and sulfhydryl donor to treat neoplastic disease)
IT
     Angiogenesis inhibitors
     Antitumor agents
     Drug interactions
     Human
        (generating angiostatin using plasminogen activator and
        sulfhydryl donor to treat neoplastic disease)
IT
     Angiogenesis
        (inhibition; generating angiostatin using plasminogen
        activator and sulfhydryl donor to treat neoplastic disease)
IT
     Neoplasm
        (metastasis; generating angiostatin using plasminogen
        activator and sulfhydryl donor to treat neoplastic disease)
IT
     354138-63-9 537677-42-2
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (angiostatin C-terminal sequence; generating angiostatin using
        plasminogen activator and sulfhydryl donor to treat neoplastic
        disease)
     337363-93-6 354138-60-6 354138-61-7
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (angiostatin N-terminal sequence; generating angiostatin using
        plasminogen activator and sulfhydryl donor to treat neoplastic
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disease)

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IT
     9001-90-5, Plasmin
                          9001-91-6, Plasminogen
     RL: BSU (Biological study, unclassified); PAC (Pharmacological activity);
     THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (angiostatin generation from; generating angiostatin using
        plasminogen activator and sulfhydryl donor to treat neoplastic
        disease)
IT
     86090-08-6, Angiostatin
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (generating angiostatin using plasminogen activator and
        sulfhydryl donor to treat neoplastic disease)
                                52-90-4, Cysteine, biological studies
TT
     52-67-5, D-Penicillamine
     70-18-8, Reduced glutathione, biological studies
                                                       616-91-1,
                        9002-01-1, Streptokinase
                                                  9039-53-6, Urokinase
     N-Acetylcysteine
     62571-86-2, Captopril
                             105913-11-9, Plasminogen activator
     139639-23-9, Tissue plasminogen activator
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
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        (generating angiostatin using plasminogen activator and
        sulfhydryl donor to treat neoplastic disease)
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                  541557-35-1
     541557-39-5
                   541557-40-8
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        angiostatin)
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     53620-20-5
                  92662-83-4
     RL: PRP (Properties)
        (unclaimed sequence; methods and compns. for generating angiostatin)
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(4) Anon; WO 9529242 1995 HCAPLUS
(5) Anon; WO 9635774 1996 HCAPLUS
(6) Anon; WO 9641194 1996 HCAPLUS
(7) Anon; WO 9715666 1997 HCAPLUS
(8) Anon; WO 9723500 1997 HCAPLUS
(9) Anon; WO 9741824 1997 HCAPLUS
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- L36 ANSWER 4 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
- AN 2003:261008 HCAPLUS
- DN 138:281097
- ED Entered STN: 04 Apr 2003
- TI Angiostatin fragments and method of use
- IN Folkman, M. Judah; O'Reilly, Michael S.; Cao, Yihai; Sim, Kim Lee
- PA USA
- SO U.S. Pat. Appl. Publ., 96 pp., Cont.-in-part of U.S. Ser. No. 335,325.

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LΑ
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IC
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NCL
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     1-6 (Pharmacology)
     Section cross-reference(s): 3, 16
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US 5792845 ECLA C12N009/68; G01N033/574
US 5885795 ECLA C12N009/68; G01N033/574
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US 6024688 ECLA C12N009/68
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 US 2004002459 ECLA
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     Fragments of an endothelial cell proliferation inhibitor and method of use
     therefor are provided. The endothelial proliferation inhibitor is a
     protein derived from plasminogen, or more specifically is an
     angiostatin fragment. The angiostatin fragments generally correspond to
     kringle structures occurring within the endothelial cell proliferation
     inhibitor. The endothelial cell inhibiting activity of these fragments
     provides a means for inhibiting angiogenesis of tumors and for
     treating angiogenic-mediated disease. Angiostatin was cloned in
     Pichia pastoris and purified from fermentation broth by lysine-Sepharose 4B.
     The purified recombinant angiostatin inhibited the bFGF-driven
     proliferation of bovine endothelial cells in vitro in a dose dependent
     manner and suppressed metastases of Lewis lung carcinoma in mice.
st
     angiostatin fragment endothelial cell proliferation inhibitor;
     angiogenesis inhibitor angiostatin fragment; antitumor angiostatin
```

fragment; metastasis Lewis lung carcinoma inhibition recombinant angiostatin TT Disease, animal (angiogenesis-mediated, treatment of; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) ΙT Bos taurus Human Macaca mulatta Mus Sus scrofa domestica (angiostatin fragment derived from plasminogen of; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) TT Angiogenesis Angiogenesis inhibitors Antiarthritics Antitumor agents Apoptosis Drug delivery systems Gene therapy Genetic vectors Mammalia Molecular cloning Protein sequences (angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) IT Blood serum (angiostatin purification from; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) IT Lung, neoplasm Mammary gland, neoplasm Prostate gland, neoplasm (carcinoma; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) Eye, disease IT (diabetic retinopathy, treatment of; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) IT Blood vessel (endothelium, cell proliferation inhibitor; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) IT (expressing angiostatin fragment; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) IT Sarcoma (fibrosarcoma; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) TT DNA Gene, animal RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (for angiostatin fragment inhibiting endothelial cell proliferation; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) Escherichia coli IT Pichia pastoris (human angiostatin expression in; angiostatin fragments as endothelial cell proliferation and angiogenesis inhibitors) IT Cell proliferation

(inhibition; angiostatin fragments as endothelial cell proliferation

and angiogenesis inhibitors)

```
Protein motifs
IT
        (kringles; angiostatin fragments as endothelial cell proliferation and
        angiogenesis inhibitors)
IT
     Eve, disease
        (macula, degeneration, treatment of; angiostatin fragments as
        endothelial cell proliferation and angiogenesis inhibitors)
IT
        (mammary; angiostatin fragments as endothelial cell proliferation and
        angiogenesis inhibitors)
IT
     Neoplasm
        (metastasis, inhibition of; angiostatin fragments as endothelial cell
        proliferation and angiogenesis inhibitors)
IT
     Lung, neoplasm
        (metastasis; angiostatin fragments as endothelial cell proliferation
        and angiogenesis inhibitors)
IT
     Transformation, genetic
        (of angiostatin fragment; angiostatin fragments as endothelial cell
       proliferation and angiogenesis inhibitors)
IT
        (prostatic; angiostatin fragments as endothelial cell proliferation and
        angiogenesis inhibitors)
IT
     Carcinoma
        (pulmonary; angiostatin fragments as endothelial cell proliferation and
        angiogenesis inhibitors)
IT
        (reticulum cell; angiostatin fragments as endothelial cell
        proliferation and angiogenesis inhibitors)
IT
     Arthritis
     Neoplasm
        (treatment of; angiostatin fragments as endothelial cell proliferation
        and angiogenesis inhibitors)
IT
     Endothelium
        (vascular, cell proliferation inhibitor; angiostatin fragments as
        endothelial cell proliferation and angiogenesis inhibitors)
IT
     506450-14-2P, Plasminogen (mouse kringle 1 fragment)
     506450-15-3P, Plasminogen (human kringle 1 fragment)
     506450-16-4P
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     fragment) 506450-18-6P, Plasminogen (cattle kringle 1
                506450-19-7P, Plasminogen (mouse kringle 2 fragment)
     fragment)
     506450-20-0P, Plasminogen (human kringle 2 fragment)
     506450-21-1P
                   506450-22-2P, Plasminogen (swine kringle 2
     fragment)
               506450-23-3P, Plasminogen (cattle kringle 2
                506450-24-4P, Plasminogen (mouse kringle 3 fragment)
     fragment)
     506450-25-5P, Plasminogen (human kringle 3 fragment)
     506450-26-6P
                    506450-27-7P, Plasminogen (swine kringle 3
     fragment)
                506450-28-8P, Plasminogen (cattle kringle 3
                 506450-29-9P, Plasminogen (mouse kringle 4 fragment)
     fragment)
     506450-30-2P, Plasminogen (human kringle 4 fragment)
     506450-31-3P, Plasminogen (mouse kringle 2-3 fragment)
     506450-32-4P, Plasminogen (human kringle 2-3 fragment)
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                   506450-34-6P, Plasminogen (swine kringle 2-3
     fragment) 506450-35-7P 506450-36-8P, Plasminogen (mouse
                            506450-37-9P, Plasminogen (human kringle
     kringle 1-3 fragment)
     1-3 fragment)
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                                   506450-39-1P, Plasminogen (swine
                            506450-40-4P
     kringle 1-3 fragment)
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     (mouse kringle 1-2 fragment)
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     kringle 1-2 fragment)
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     (swine kringle 1-2 fragment)
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     Plasminogen (mouse kringle 1-4 fragment)
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     Plasminogen (human kringle 1-4 fragment)
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     Plasminogen (mouse kringle 1-4BKLS)
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     Plasminogen (human kringle 1-4BKLS)
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RL: BPN (Biosynthetic preparation); BSU (Biological study, unclassified);

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PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use);
     BIOL (Biological study); PREP (Preparation); USES (Uses)
        (amino acid sequence, as angiostatin fragment; angiostatin fragments as
        endothelial cell proliferation and angiogenesis inhibitors)
     506450-13-1, Angiostatin (Rhesus monkey)
IT
                                               506450-50-6,
     Plasminogen (mouse)
                          506450-51-7, Angiostatin (mouse)
                                       506450-53-9, Angiostatin (swine)
     506450-52-8, Angiostatin (human)
     506450-54-0, Angiostatin (cattle)
     RL: BSU (Biological study, unclassified); PRP (Properties); BIOL
     (Biological study)
        (amino acid sequence; angiostatin fragments as endothelial cell
       proliferation and angiogenesis inhibitors)
IT
     9001-91-6, Plasminogen
     RL: BSU (Biological study, unclassified); RCT (Reactant); BIOL (Biological
     study); RACT (Reactant or reagent)
        (angiostatin fragment derived from; angiostatin fragments as
        endothelial cell proliferation and angiogenesis inhibitors)
IT
     86090-08-6P, Angiostatin
     RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BSU
     (Biological study, unclassified); PAC (Pharmacological activity); PUR
     (Purification or recovery); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (angiostatin fragments as endothelial cell proliferation and
        angiogenesis inhibitors)
     9012-36-6D, Sepharose 4B, conjugates with lysine
IT
     RL: NUU (Other use, unclassified); USES (Uses)
        (in angiostatin purification; angiostatin fragments as endothelial cell
       proliferation and angiogenesis inhibitors)
IT
     506457-30-3
                  506457-31-4
                               506457-32-5
     RL: PRP (Properties)
        (unclaimed nucleotide sequence; angiostatin fragments and method of
       use)
IT
     122580-21-6
                  506457-33-6
     RL: PRP (Properties)
        (unclaimed sequence; angiostatin fragments and method of use)
    ANSWER 5 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
L36
AN
     2000:283962 HCAPLUS
DN
     132:304929
ED
     Entered STN: 03 May 2000
     Method of making mammalian kringle 5 peptide fragments with
TT
     angiogenesis inhibitory effect by elastase proteolytic cleavage of
     plasminogen
IN
    Davidson, Donald J.
     Abbott Laboratories, USA
PA
SO
     U.S., 48 pp., Cont.-in-part of U.S. Ser. No. 832,087.
     CODEN: USXXAM
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     435068100
     6-3 (General Biochemistry)
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A2

PRAI US 1996-643219

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robinson - 10 / 753646
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US 5981484 ECLA C12N009/68
US 6699838 ECLA C12N009/68
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US 2004138127 ECLA C12N009/68
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    A method of making mammalian kringle 5 peptide fragments corresponding to
    the 5th kringle domain of mammalian plasminogen and having
     angiogenic inhibitory effect is claimed. The method comprises
     exposing a mammalian plasminogen to elastase at a ratio of about
     1:100 to 1:300 (weight/weight) and isolating kringle 5 fragments from the
mixture
    Kringle 5 peptide fragments were prepared either by porcine elastase
    proteolytic cleavage of Lys plasminogen or synthesized by standard
     solid phase Fmoc chemical The inhibition of bovine capillary endothelial
     cell proliferation and migration by kringle 5 peptide fragments was both
    potent and specific to the endothelial cells but not normal or tumor
     cells. Kringle 5 peptide fragments were also produced recombinantly in
     Pichia pastoris and E. coli.
ST
     elastase cleavage mammalian plasminogen kringle 5 peptide
     isolation; kringle 5 peptide sequence mammalian plasminogen
     angiogenesis inhibition
```

ΙT Angiogenic factors

Angiogenic factors

Growth inhibitors, animal

Growth inhibitors, animal

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); PROC (Process)

(angiogenic growth-inhibiting

factors; method of making mammalian kringle 5 peptide fragments with angiogenesis inhibitory effect by elastase proteolytic cleavage of plasminogen)

IT Blood vessel

> (endothelium, proliferation and migration of, inhibition by kringle 5 peptide fragments; method of making mammalian kringle 5 peptide fragments with angiogenesis inhibitory effect by elastase proteolytic cleavage of plasminogen)

Escherichia coli IT

Komagataella pastoris

(expression host; method of making mammalian kringle 5 peptide fragments with angiogenesis inhibitory effect by elastase proteolytic cleavage of plasminogen)

IT Cell migration

> (inhibitors, of bovine capillary endothelial cell; method of making mammalian kringle 5 peptide fragments with angiogenesis inhibitory effect by elastase proteolytic cleavage of plasminogen)

ΙT Protein motifs

> (kringles; method of making mammalian kringle 5 peptide fragments with angiogenesis inhibitory effect by elastase proteolytic cleavage of plasminogen)

. IT Peptides, biological studies

RL: BAC (Biological activity or effector, except adverse); BPR (Biological

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process); BSU (Biological study, unclassified); PRP (Properties); SPN
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     (Process)
        (method of making mammalian kringle 5 peptide fragments with
        angiogenesis inhibitory effect by elastase proteolytic cleavage
        of plasminogen)
IT
     Cytotoxic agents
        (of bovine capillary endothelial cell; method of making mammalian
        kringle 5 peptide fragments with angiogenesis inhibitory
        effect by elastase proteolytic cleavage of plasminogen)
     Protein sequences
TT
        (of human plasminogen fragments; method of making mammalian
        kringle 5 peptide fragments with angiogenesis inhibitory
        effect by elastase proteolytic cleavage of plasminogen)
     Proliferation inhibition
IT
        (proliferation inhibitors, of bovine capillary endothelial cell; method
        of making mammalian kringle 5 peptide fragments with
        angiogenesis inhibitory effect by elastase proteolytic cleavage
        of plasminogen)
     9001-91-6D, Lys plasminogen, de-(1-76) derivs.
TΤ
     RL: BPR (Biological process); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); PROC (Process)
        (Lys plasminogen, kringle 5 peptide fragment source; method
        of making mammalian kringle 5 peptide fragments with
        angiogenesis inhibitory effect by elastase proteolytic cleavage
        of plasminogen)
IT
     265110-89-2
     RL: PRP (Properties)
        (Unclaimed; method of making mammalian kringle 5 peptide fragments with
        angiogenesis inhibitory effect by elastase proteolytic cleavage
        of plasminogen)
IT
     250159-78-5D, 443-543-Plasminogen (human), peptides
     250159-79-6D, 449-543-Plasminogen (human), peptides
     250159-80-9D, 454-543-Plasminogen (human), peptides
     250159-81-0D, 443-546-Plasminogen (human), peptides
     250159-83-2D, 449-546-Plasminogen (human), peptides
     250159-84-3D, 454-546-Plasminogen (human), peptides
     264868-26-0D, 355-543-Plasminogen (human), peptides
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     (Biological study); PROC (Process)
        (amino acid sequence; method of making mammalian kringle 5 peptide
        fragments with angiogenesis inhibitory effect by elastase
        proteolytic cleavage of plasminogen)
IT
     109884-31-3, Plasminogen (human)
     RL: BPR (Biological process); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); PROC (Process)
        (amino acid sequence; method of making mammalian kringle 5 peptide
        fragments with angiogenesis inhibitory effect by elastase
        proteolytic cleavage of plasminogen)
IT
     9001-91-6, Plasminogen
     RL: BPR (Biological process); BSU (Biological study, unclassified); PRP
     (Properties); BIOL (Biological study); PROC (Process)
        (method of making mammalian kringle 5 peptide fragments with
        angiogenesis inhibitory effect by elastase proteolytic cleavage
        of plasminogen)
IT
     9004-06-2, Elastase
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (method of making mammalian kringle 5 peptide fragments with
        angiogenesis inhibitory effect by elastase proteolytic cleavage
        of plasminogen)
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IT
     199664-76-1P
                    199664-77-2P
                                                  199664-81-8P
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IT

TΤ

RE

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     199664-88-5P 199664-89-6P 199664-90-9P
     199664-91-0P
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PRP (Properties); SPN (Synthetic preparation); BIOL
     (Biological study); PREP (Preparation)
        (synthesis of, antiangiogenic kringle 5 peptide; method of
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        inhibitory effect by elastase proteolytic cleavage of
       plasminogen)
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        elastase proteolytic cleavage of plasminogen)
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(1) Anon; WO 9204450 1992 HCAPLUS
(2) Anon; WO 9529242 1995 HCAPLUS
(3) Anon; WO 9723500 1997 HCAPLUS
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   MEDLINE
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   V3, P191 HCAPLUS
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      P97 HCAPLUS
(29) Weidner, N; The New England Journal of Medicine 1991, V324(1), P1 MEDLINE
L36 ANSWER 6 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
        1999:764061 HCAPLUS
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        Entered STN: 03 Dec 1999
ED
        Preparation of peptides as anti-angiogenic drugs to treat cancer,
ΤI
        arthritis and retinopathy
IN
        Kawai, Megumi; Henkin, Jack; Sheppard, George S.; Craig, Richard A.
PA
        Abbott Laboratories, USA
SO
        PCT Int. Appl., 34 pp.
        CODEN: PIXXD2
DT
        Patent
T.A
        English
        ICM C07K005-11
IC
        ICS C07K005-065
        34-3 (Amino Acids, Peptides, and Proteins)
        Section cross-reference(s): 1
FAN.CNT 1
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                                                                        APPLICATION NO.
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        WO 9961466
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 WO 9961466
AB
        Peptides WNR1CR2R3CRARBNXCR4R5CRCRDNYCR6R7CRERFNZCR8R9CRGRHR10 [RARB,
        RCRD, RERF, or RGRH = H or :O; W, X, Y, Z = H, alkyl; R1 = H, protective
        group; R2, R3 = H, aminoalkyl; R4, R5 = H, alkyl, cycloalkyl; R6, R7 = H,
        alkyl, arylalkyl; R8, R9 = H, alkyl, carboxy- or carbalkoxyalkyl; R10 = OH, (un)substituted alkoxy, cycloalkoxy, NH2, (un)substituted alkylamino,
        cycloalkylamino] were prepared for treating pathol. states which arise from
        or are exacerbated by angiogenesis. Thus, (2S)-2-[[(2S)-2-[[(2S)-2-[[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-[(2S)-2-
        2-(acetylamino)-6-aminohexanoyl]amino]-4-methylpentanoyl]methylamino]-3-
        phenylpropanoyl]amino]butanedioic acid was prepared and showed 83%
        inhibition at 10 NM against human microvascular endothelial cell
        migration.
ST
        peptide prepn angiogenesis inhibitor; antitumor treatment peptide prepn;
        antiarthritic peptide prepn; retinopathy treatment peptide prepn
IT
        Angiogenesis inhibitors
        Antiarthritics
        Antitumor agents
              (preparation of peptides as anti-angiogenic drugs to treat cancer, arthritis
              and retinopathy)
IT
        Peptides, preparation
        RL: BAC (Biological activity or effector, except adverse); BSU (Biological
        study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
        BIOL (Biological study); PREP (Preparation); USES (Uses)
              (preparation of peptides as anti-angiogenic drugs to treat cancer, arthritis
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and retinopathy)

Eye, disease IT (retinopathy; preparation of peptides as anti-angiogenic drugs to treat cancer, arthritis and retinopathy) TΤ 251555-82-5P 251555-83-6P 251555-84-7P 251555-85-8P 251555-86-9P RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (preparation of peptides as anti-angiogenic drugs to treat cancer, arthritis and retinopathy) 251555-87-0P 251555-88-1P 251555-89-2P 251555-90-5P TΤ 251555-91-6P 251555-92-7P 251555-93-8P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of peptides as anti-angiogenic drugs to treat cancer, arthritis and retinopathy) RE.CNT THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD RE (1) Abbott Lab; WO 9741824 A 1997 HCAPLUS (2) Childrens Medical Center; WO 9529242 A 1995 HCAPLUS (3) Eyal, J; US 5654277 A 1997 HCAPLUS (4) Grace W R & Co; EP 0514721 A 1992 HCAPLUS (5) Weinstein, B; Chemistry and Biochemistry of Amino Acids Peptides and Proteins 1983, V7, P266 L36 ANSWER 7 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN AN 1999:718961 HCAPLUS DN 131:346531 ED Entered STN: 11 Nov 1999 ΤI antiangiogenic kringle 5 peptide fragments of plasminogen for therapeutic control of angiogenesis IN Davidson, Donald J. PA Abbott Laboratories, USA SO U.S., 40 pp., Cont.-in-part of U.S. 5,801,146. CODEN: USXXAM DTPatent LA English IC ICM A61K038-00 ICS A61K038-04 NCL 514012000 1-8 (Pharmacology) Section cross-reference(s): 14 FAN.CNT 4 PATENT NO. KIND DATE APPLICATION NO. DATE ---------------_____ US 5981484 À 19991109 US 1997-832087 19970403 <--US 5801146 Α 19980901 US 1996-643219 19960503 <--EP 910571 A2 19990428 EP 1997-925478 19970505 <--R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, FI CN 1223690 Α 19990721 CN 1997-195989 19970505 <--BR 9708911 Α 19990803 BR 1997-8911 19970505 <--US 6057122 US 1997-851350 Α 20000502 19970505 <--NZ 1997-332319 NZ 332319 Α 20000929 19970505 <--JP 2002502235 T2 20020122 JP 1997-540162 19970505 <--US 6699838 B1 20040302 US 1997-924287 19970905 <--US 5972896 Α 19991026 US 1998-131995 19980811 <--US 6251867 В1 20010626 US 1998-132154 19980811 <--KR 2000010739 Α 20000225 KR 1998-708851 19981103 <--

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    Mammalian kringle 5 peptide fragments that can inhibit
     angiogenesis are described for treating angiogenic
     diseases. Kringle 5 peptide fragments were manufactured either by proteolytic
     cleavage of plasminogens from various species or synthesized by
     standard FMOC chemical The inhibition of stimulated proliferation and
migration
    by kringle 5 peptide fragments was both potent and specific to the bovine
     endothelial cells but not normal or tumor cells. Methods and compns. for
     inhibiting angiogenic diseases are also proposed.
st
    kringle 5 peptide plasminogen angiogenesis treatment;
     antiangiogenic kringle 5 peptide plasminogen
IT
     Angiogenesis inhibitors
        (antiangiogenic kringle 5 peptide fragments of
       plasminogen for therapeutic control of angiogenesis)
IT
    Antiarthritics
    Antitumor agents
        (antiangiogenic plasminogen kringle 5 domain
       peptides as, for inhibition of angiogenesis;
       antiangiogenic kringle 5 peptide fragments of
       plasminogen for therapeutic control of angiogenesis)
ΙT
    Eye, disease
        (diabetic retinopathy, treatment of; antiangiogenic kringle 5
       peptide fragments of plasminogen for therapeutic control of
       angiogenesis)
IT
    Blood vessel
        (endothelium, proliferation inhibition by human kringle 5 peptide;
       antiangiogenic kringle 5 peptide fragments of
       plasminogen for therapeutic control of angiogenesis)
IT
    Protein sequences
        (for plasminogen of human; antiangiogenic kringle 5
       peptide fragments of plasminogen for therapeutic control of
       angiogenesis)
IT
    Blood vessel, neoplasm
    Blood vessel, neoplasm
        (hemangioma, inhibitors; antiangiogenic kringle 5 peptide
       fragments of plasminogen for therapeutic control of
       angiogenesis)
IT
    Antitumor agents
        (hemangioma; antiangiogenic kringle 5 peptide fragments of
       plasminogen for therapeutic control of angiogenesis)
IT
    Protein motifs
        (kringles, fragments of, human plasminogen
       angiogenesis inhibitors; antiangiogenic kringle 5
       peptide fragments of plasminogen for therapeutic control of
       angiogenesis)
IT
    Antitumor agents
        (lymphoma; antiangiogenic kringle 5 peptide fragments of
       plasminogen for therapeutic control of angiogenesis)
IT
    Eye, disease
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(macula, degeneration, treatment of; antiangiogenic kringle 5
        peptide fragments of plasminogen for therapeutic control of
        angiogenesis)
IT
     Cattle
     Macaca mulatta
     Mouse
     Swine
        (plasminogen kringle 5 domain peptides of, for inhibition of
        angiogenesis; antiangiogenic kringle 5 peptide
        fragments of plasminogen for therapeutic control of
        angiogenesis)
IT
     Antitumor agents
        (sarcoma; antiangiogenic kringle 5 peptide fragments of
        plasminogen for therapeutic control of angiogenesis)
     Psoriasis
TT
        (treatment of; antiangiogenic kringle 5 peptide fragments of
        plasminogen for therapeutic control of angiogenesis)
IT
     109884-31-3D, Plasminogen (human liver clone pPLGKG protein
     moiety reduced), peptides 250159-78-5D, 443-543-
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     RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES
        (amino acid sequence; antiangiogenic kringle 5 peptide
        fragments of plasminogen for therapeutic control of
        angiogenesis)
IΤ
     9001-91-6, Plasminogen
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (antiangiogenic peptides of; antiangiogenic kringle
        5 peptide fragments of plasminogen for therapeutic control of
        angiogenesis)
     199664-76-1P
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TΤ
                    199664-77-2P
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     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); SPN (Synthetic preparation); BIOL (Biological
     study); PREP (Preparation)
        (synthesis of, antiangiogenic kringle 5 peptide;
        antiangiogenic kringle 5 peptide fragments of
        plasminogen for therapeutic control of angiogenesis)
     140088-37-5
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     RL: PRP (Properties)
        (unclaimed nucleotide sequence; antiangiogenic kringle 5
        peptide fragments of plasminogen for therapeutic control of
        angiogenesis)
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                                 250163-85-0 250163-86-1
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        (unclaimed protein sequence; antiangiogenic kringle 5 peptide
        fragments of plasminogen for therapeutic control of
        angiogenesis)
RE.CNT
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              THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD
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- L36 ANSWER 8 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
- AN 1997:740418 HCAPLUS
- DN 128:43873
- ED Entered STN: 24 Nov 1997
- TI Antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis
- IN Davidson, Donald J.; Wang, Jieyi; Gubbins, Earl J.
- PA Abbott Laboratories, USA

US 1997-832087 A

- SO PCT Int. Appl., 78 pp. CODEN: PIXXD2
- DT Patent
- LA English
- IC ICM A61K
- CC 1-12 (Pharmacology)

FAN.CNT 4

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    Mammalian kringle 5 fragments and kringle 5 fusion proteins are disclosed
AB
     as compds. for treating angiogenic diseases. Methods and
     compns. for inhibiting angiogenic diseases are also disclosed.
ST
    kringle 5 peptide antiangiogenesis sequence
TT
    Angiogenic factors
      Angiogenic factors
      Growth inhibitors, animal
      Growth inhibitors, animal
     RL: PEP (Physical, engineering or chemical process); PRP (Properties); THU
     (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (angiogenic growth-inhibiting
        factors; antiangiogenic peptides, polypeptides containing
        them, and methods for inhibiting angiogenesis)
    Angiogenesis inhibitors
IT
    Antiarthritics
    Antitumor agents
     Escherichia coli
    Gene therapy
    Genetic vectors
     Komagataella pastoris
     Protein sequences
    RNA sequences
     cDNA sequences
        (antiangiogenic peptides, polypeptides containing them, and
       methods for inhibiting angiogenesis)
IT
     Fusion proteins (chimeric proteins)
     RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic
     use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (antiangiogenic peptides, polypeptides containing them, and
       methods for inhibiting angiogenesis)
IT
    Antitumor agents
        (carcinoma; antiangiogenic peptides, polypeptides containing
       them, and methods for inhibiting angiogenesis)
TT
     Eye, disease
        (diabetic retinopathy, inhibitors; antiangiogenic peptides,
       polypeptides containing them, and methods for inhibiting
       angiogenesis)
IT
    Blood vessel
        (endothelium, migration of cells of; antiangiogenic peptides,
       polypeptides containing them, and methods for inhibiting
       angiogenesis)
TΤ
    Blood vessel, neoplasm
        (hemangioma, inhibitors; antiangiogenic peptides,
       polypeptides containing them, and methods for inhibiting
       angiogenesis)
IT
     Psoriasis
        (inhibitors; antiangiogenic peptides, polypeptides containing
        them, and methods for inhibiting angiogenesis)
IT
    Cattle
    Macaca mulatta
    Mouse
     Swine
        (kringle 5 fusion protein of; antiangiogenic peptides,
       polypeptides containing them, and methods for inhibiting
       angiogenesis)
TΤ
    Protein motifs
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(kringles; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT Antitumor agents (lymphoma; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) TT Eye, disease (macula, degeneration, inhibitors; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) Antitumor agents IT (metastasis; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT Cell migration (of vascular endothelium; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT Antitumor agents (sarcoma; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) 109884-31-3, Plasminogen (human liver clone pPLGKG protein IT moiety reduced) RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study); OCCU (Occurrence) (amino acid sequence; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT 29022-11-5P 35661-39-3P 35661-40-6P 71989-26-9P 71989-28-1P 119831-72-0P 132388-59-1P 199664-76-1P 199664-77-2P 199664-80-7P 199664-81-8P 199664-82-9P 199664-83-0P 199664-84-1P 199664-85-2P 199664-86-3P 199664-87-4P 199664-88-5P 199664-89-6P 199664-90-9P 199664-91-0P RL: PNU (Preparation, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT 35661-60-0 35737-15-6 68858-20-8 71989-14-5 71989-18-9 71989-23-6 71989-31-6 71989-33-8 71989-35-0 109425-51-6 RL: RCT (Reactant); RACT (Reactant or reagent) (antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT 9001-91-6, Plasminogen RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process) (elastase treatment of; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT 56-84-8, L-Aspartic acid, biological studies 56-87-1, Lysine, biological 60-18-4, Tyrosine, biological studies 61-90-5, L-Leucine, biological studies 63-91-2, Phenylalanine, biological studies L-Arginine, biological studies RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence) (kringle peptide containing; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) IT 9004-06-2, Elastase RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study) (plasminogen treatment with; antiangiogenic peptides, polypeptides containing them, and methods for inhibiting angiogenesis) L36 ANSWER 9 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN

1997:595457 HCAPLUS

AN

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DN
     127:304391
ED
     Entered STN: 18 Sep 1997
     Kringle 5 of plasminogen is a novel inhibitor of endothelial
TΤ
     cell growth
     Cao, Yihai; Chen, Andrew; An, Seong Soo A.; Ji, Richard-Weidong;
AU
     Davidson, Don; Cao, Yumei; Llinas, Miguel
     Laboratory of Angiogenesis Research, Department of Cell and Molecular
CS
     Biology, Karolinska Institute, Stockholm, S-17177, Swed.
     Journal of Biological Chemistry (1997), 272(36), 22924-22928
so
     CODEN: JBCHA3; ISSN: 0021-9258
PB
     American Society for Biochemistry and Molecular Biology
DT
     Journal
LΑ
     English
CC
     6-3 (General Biochemistry)
     Section cross-reference(s): 13
     Angiostatin is a potent angiogenesis inhibitor which has been identified
AB
     as an internal fragment of plasminogen that includes its first
     four kringle modules. We have recently demonstrated that the
     anti-endothelial cell proliferative activity of angiostatin is also
     displayed by the first three kringle structures of plasminogen
     and marginally so by kringle 4 (Cao, Y., Ji, R.-W., Davidson, D.,
     Schaller, J., Marti, D., Sohndel, S., McCance, S. G., O'Reilly, M. S.,
     Llinas, M., and Folkman, J. (1996) J. Biol. Chemical 271, 29461-29467).
     now report that the kringle 5 fragment of human plasminogen is a
     specific inhibitor for endothelial cell proliferation. Kringle 5 obtained
     as a proteolytic fragment of human plasminogen displays potent
     inhibitory effect on bovine capillary endothelial cells with a
     half-maximal concentration (ED50) of approx. 50 nM. Thus, kringle 5 would
appear
     to be more potent than angiostatin on inhibition of basic fibroblast
     growth factor-stimulated capillary endothelial cell proliferation.
     Appropriately folded recombinant mouse kringle 5 protein, expressed in
     Escherichia coli, exhibits a comparable inhibitory effect as the
     proteolytic kringle 5 fragment. Thus, kringle 5 domain of human
    plasminogen is a novel endothelial inhibitor that is sufficiently
     potent to block the growth factor-stimulated endothelial cell growth.
ST
     kringle domain plasminogen endothelial cell growth
     Blood vessel
TΤ
        (endothelium; kringle 5 of plasminogen is a novel inhibitor
        of endothelial cell growth)
IT
     Protein sequences
        (kringle 5 of plasminogen is a novel inhibitor of endothelial
        cell growth)
IT
     Protein motifs
        (kringles; kringle 5 of plasminogen is a novel inhibitor of
        endothelial cell growth)
IT
     196417-08-0
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); PRP (Properties); BIOL (Biological study)
        (amino acid sequence; kringle 5 of plasminogen is a novel
        inhibitor of endothelial cell growth)
IT
     9001-91-6, Plasminogen
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); BIOL (Biological study)
        (kringle 5 of plasminogen is a novel inhibitor of endothelial
        cell growth)
RE.CNT
              THERE ARE 39 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Brooks, P; Cell 1994, V79, P1157 HCAPLUS
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(6) Castellino, F; Methods Enzymol 1981, V80, P365 HCAPLUS (7) Chen, C; Cancer Res 1995, V55, P4230 HCAPLUS (8) Clapp, C; Endocrinology 1993, V133, P1292 HCAPLUS (9) Cox, M; Chem Phys Lipids 1994, V67, P43 (10) Dameron, K; Science 1994, V265, P1582 HCAPLUS (11) Deutsch, D; Science 1970, V170, P1095 HCAPLUS (12) Dong, Z; Cell 1997, V88, P801 HCAPLUS (13) Ferrara, N; Endocr Rev 1992, V13, P18 HCAPLUS (14) Folkman, J; Cell 1996, V87, P1153 HCAPLUS (15) Folkman, J; J Biol Chem 1992, V267, P10931 HCAPLUS (16) Folkman, J; N Engl J Med 1995, V333, P1757 MEDLINE (17) Folkman, J; Nat Med 1995, V1, P27 HCAPLUS (18) Folkman, J; Proc Natl Acad Sci U S A 1979, V76, P5217 MEDLINE (19) Friedlander, M; Science 1995, V270, P1500 HCAPLUS (20) Gately, S; Cancer Res 1996, V56, P4887 HCAPLUS (21) Good, D; Proc Natl Acad Sci U S A 1990, V87, P6624 HCAPLUS (22) Grant, D; Cell 1989, V58, P933 HCAPLUS (23) Gupta, S; Proc Natl Acad Sci U S A 1995, V92, P7799 HCAPLUS (24) Hanahan, D; Cell 1996, V86, P353 HCAPLUS (25) Hartree, E; Anal Biochem 1972, V48, P422 HCAPLUS (26) Homandberg, G; Am J Pathol 1985, V120, P327 HCAPLUS (27) Hori, A; Cancer Res 1991, V51, P6180 HCAPLUS (28) Kandel, J; Cell 1991, V66, P1095 HCAPLUS (29) Menhart, N; Biochemistry 1993, V32, P8799 HCAPLUS (30) Mulichak, A; Biochemistry 1991, V30, P10576 HCAPLUS (31) Nguyen, M; J Natl Cancer Inst 1994, V86, P356 MEDLINE (32) O'Reilly, M; Cell 1994, V79, P315 HCAPLUS (33) O'Reilly, M; Cell 1997, V88, P1 (34) O'Reilly, M; Nat Med 1996, V2, P689 HCAPLUS (35) Sage, E; J Cell Biochem 1995, V57, P127 HCAPLUS (36) Senger, D; Cancer Res 1990, V50, P1774 HCAPLUS (37) Sottrup-Jensen, L; Prog Chem Fibrinolysis Thrombolysis 1978, V3, P191 **HCAPLUS** (38) Thewes, T; Biochem Biophys Acta 1987, V912, P254 HCAPLUS (39) Tolsma, S; J Cell Biol 1993, V122, P497 HCAPLUS L36 ANSWER 10 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN AN1997:48745 HCAPLUS DN 126:54861 ED Entered STN: 23 Jan 1997 TI Angiostatin krinkle region-containing fragment sequences, aggregate angiostatin, and tumor or angiogenic-mediated disease treatment Folkman, M. Judah; O'Reilly, Michael S.; Cao, Yihai; Sim, Kim Lee; Lin, IN PΑ Children's Medical Center Corporation, USA SÓ PCT Int. Appl., 212 pp. CODEN: PIXXD2 DT Patent LA English IC ICM C12N CC 1-6 (Pharmacology) Section cross-reference(s): 6, 13 FAN.CNT 6 PATENT NO. KIND DATE APPLICATION NO. DATE -**-**----_ _ _ _ _ _ _ _ _ _ _ _ ------WO 1996-US5856 PΙ WO 9635774 A2 19961114 19960426 <--WO 9635774 Α3 19970213 AL, AM, AT, AU, AZ, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IS, JP, KE, KG, KP, KR, KZ, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR,

IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML

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 WO 9635774 ICM
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 US 5885795
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US 5837682 ECLA C12N009/68
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     Fragments and an aggregate form of an endothelial cell proliferation
     inhibitor and methods of use therefor are provided. The endothelial
     proliferation inhibitor is a protein from plasminogen, or more
     specifically is an angiostatin fragment. The angiostatin fragments
     generally correspond to krinkle structures occurring within the
     endothelial cell proliferation inhibitor. Angiostatin is also prepared in a
     aggregate form. The endothelial cell inhibiting activity of the
     angiostatin fragments and the aggregate angiostatin provide a means for
     inhibiting angiogenesis of tumors and for treating
     angiogenic-mediated diseases.
ST
     angiostatin krinkle region sequence disease treatment;
     angiogenesis inhibitor angiostatin krinkle region sequence;
     aggregate angiostatin krinkle region disease treatment; tumor
     angiogenesis inhibition angiostatin krinkle region;
     plasminogen angiostatin fragment sequence disease treatment
IT
     Antitumor agents
     Cattle
     Macaca mulatta
     Mouse
     Protein sequences
        (angiostatin krinkle region-containing fragment sequences, aggregate
        angiostatin, and tumor or angiogenic-mediated disease
        treatment)
IT
     Apoptosis
        (angiostatin-stimulated; angiostatin krinkle region-containing fragment
        sequences, aggregate angiostatin, and tumor or angiogenic
        -mediated disease treatment)
IT
     Lung, neoplasm
        (carcinoma, treatment; angiostatin krinkle region-containing fragment
        sequences, aggregate angiostatin, and tumor or angiogenic
        -mediated disease treatment)
```

IT Escherichia coli

Komagataella pastoris

(expression host; angiostatin krinkle region-containing fragment sequences, aggregate angiostatin, and tumor or **angiogenic**-mediated disease treatment)

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IT
    Lung, neoplasm
        (inhibitors; angiostatin krinkle region-containing fragment sequences,
        aggregate angiostatin, and tumor or angiogenic-mediated
        disease treatment)
     Antitumor agents
IT
        (lung; angiostatin krinkle region-containing fragment sequences, aggregate
        angiostatin, and tumor or angiogenic-mediated disease
        treatment)
     122071-87-8, 84-162-Plasminogen (human liver clone
IT
     pPLGKG protein moiety reduced) 185074-38-8
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     RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (amino acid sequence; angiostatin krinkle region-containing fragment
        sequences, aggregate angiostatin, and tumor or angiogenic
        -mediated disease treatment)
                              86090-08-6, Angiostatin
IT
     9001-91-6, Plasminogen
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     172642-30-7, Angiostatin (human)
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               172642-32-9, Angiostatin (pig)
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     RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (angiostatin krinkle region-containing fragment sequences, aggregate
        angiostatin, and tumor or angiogenic-mediated disease
        treatment)
    ANSWER 11 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
L36
     1995:842649 HCAPLUS
ΑN
DN
     123:246823
ED
     Entered STN: 10 Oct 1995
ΤI
    Hydrophilic signal oligopeptides and methods of therapeutic use
IN
     Rath, Matthias
PA
SO
     PCT Int. Appl., 87 pp.
     CODEN: PIXXD2
DТ
     Patent
LΑ
    English
TC
     ICM G01N033-531
CC
     1-7 (Pharmacology)
     Section cross-reference(s): 6, 7, 15
FAN.CNT 1
     PATENT NO.
                         KIND
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                                            APPLICATION NO.
                                                                   DATE
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PΤ
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                         A1
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            MX, NL, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TT, UA, US,
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US 2005014138

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20050120

US 2004-930300

20040830 <--

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US 2001-881976
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CLASS
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CLASS PATENT FAMILY CLASSIFICATION CODES PATENT NO.

ICM G01N033-531 WO 9519568

The instant invention is directed to a method of identifying signal oligopeptides through the use of algorithms, the use of signal oligopeptides as vaccines and as immunogens to produce antibodies. Like the human language, the protein code consists of letters, words, and sentences. The letters (amino acids) and sentences (complete 3-dimensional proteins) have been known previously, but the present discovery identifies the protein words or verbs. These protein verbs are represented by signal oligopeptides which are localized on the surface of the protein and are represented by the hydrophilicity maxima of the protein. These signal oligopeptides are enriched in charged amino acids in a versatile arrangement with neutral spacer amino acids. The sp. signal character of these oligopeptides is determined by a characteristic combination of conformation and charge within the signal sequence. Sas in human language, the whole sentence (complete 3-dimensional protein) is needed to determine the sp. and complete action of any given protein. language eliminating or changing the verb of a sentence renders the whole sentence meaningless. Similarly, blocking the protein code verbs (signal oligopeptides) can be therapeutically used to block the undesired action or interaction of an entire protein. The discovery of the protein code provides the rationale for deciphering the communication code of diseases. Infectious diseases, cancer, cardiovascular and other diseases develop by means of one or more pathogenicity-mediating protein. Blocking the signal oligopeptides of these proteins (e.g., with antibodies) allows the sp. therapeutic interception of a pathol. communication and thereby blocks disease propagation. Some 360 oligopeptides of signal significance are presented.

hydrophilic signal oligopeptide code sequence therapy; antibody signal ST oligopeptide sequence therapy

Proteins, specific or class TT

RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)

(ACTH-releasing factor-binding, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use)

IT Schistosoma

> (elastase precursor of; hydrophilic signal oligopeptides and methods of therapeutic use)

IT Algorithm

> (for signal peptide searching; hydrophilic signal oligopeptides and methods of therapeutic use)

IT Antibodies

IT

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (hydrophilic signal oligopeptide-binding; hydrophilic signal oligopeptides and methods of therapeutic use)

TТ Acquired immune deficiency syndrome

Hydrophilicity

Therapeutics

(hydrophilic signal oligopeptides and methods of therapeutic use) Treponema pallidum

(membrane protein TMPA of, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use)

Proteins, properties IT

RL: PRP (Properties)

(protein functional code; hydrophilic signal oligopeptides and methods

of therapeutic use) TΤ Hepatitis (δ antigen; hydrophilic signal oligopeptides and methods of therapeutic use) Mental disorder IT (Alzheimer's disease, amyloid A4; hydrophilic signal oligopeptides and methods of therapeutic use) Glycoproteins, specific or class IT RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (B, of herpes virus 1, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) ΙT Lipoproteins RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (Lp(a), apo-, human and rhesus, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) Antigens, IT RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (TmpA (treponemal membrane protein A), of Treponema pallidum, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) ITProteins, specific or class RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (amyloid A4, Alzheimer, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) IT Lipoproteins RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (apo-, E, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) TТ Phosphoproteins RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (gene rev, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) TΤ Virus, animal RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (herpes simplex 1, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) Virus, animal RL: BPR (Biological process); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (herpes simplex 2, glycoprotein B of, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) ITPeptides, biological studies RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (oligo-, hydrophilic signal oligopeptides and methods of therapeutic use)

9001-12-1, Collagenase RL: BPR (Biological process); BSU (Biological study, unclassified); PRP

IT

(Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (fibroblast MMP1, signal fragments; hydrophilic signal oligopeptides and methods of therapeutic use) IT 168690-10-6 168690-11-7 99713-67-4 168690-08-2 168690-09-3 168690-16-2 168690-12-8 168690-13-9 168690-14-0 168690-15-1 . 168690-21-9 168690-17-3 168690-18-4 168690-19-5 168690-20-8 168690-26-4 168690-22-0 168690-23-1 168690-24-2 168690-25-3 168690-31-1 168690-27-5 168690-28-6 168690-29-7 168690-30-0 168690-32-2 168690-33-3 168690-34-4 168690-35-5 168690-36-6 168690-37-7 168690-38-8 168690-39-9 168690-40-2 168690-41-3 168690-42-4 168690-43-5 168690-44-6 168690-45-7 168690-46-8 168690-47-9 168690-48-0 168690-49-1 168690-50-4 168690-51-5 168690-52-6 168690-53-7 168690-54-8 168690-55-9 168690-56-0 168690-57-1 168690-58-2 168690-59-3 168690-60-6 168690-61-7 168690-62-8 168690-63-9 168690-64-0 168690-65-1 168690-66-2 168690-68-4 168690-69-5 168690-70-8 168690-71-9 168690-67-3 168690-72-0 168690-73-1 168690-74-2 168690-75-3 168690-76-4 168690-77-5 168690-78-6 168690-79-7 168690-80-0 168690-81-1 168690-83-3 168690-84-4 168690-85-5 168690-86-6 168690-82-2 168690-87-7 168690-88-8 168690-89-9 168690-90-2 168690-91-3 168690-92-4 168690-93-5 168690-94-6 168690-95-7 168690-96-8 168690-97-9 168690-98-0 168690-99-1 168691-00-7 168691-01-8 168691-02-9 168691-03-0 168691-04-1 168691-05-2 168691-06-3 168691-07-4 168691-08-5 168691-09-6 168691-10-9 168691-11-0 168691-13-2 168691-14-3 168691-15-4 168691-16-5 168691-12-1 168691-18-7 168691-19-8 168691-20-1 168691-21-2 168691-17-6 168691-22-3 168691-23-4 168691-24-5 168691-25-6 168691-26-7 168691-27-8 168691-28-9 168691-29-0 168691-30-3 168691-31-4 168691-34-7 168691-35-8 168691-36-9 168691-32-5 168691-33-6 168691-39-2 168691-40-5 168691-41-6 168691-37-0 168691-38-1 168691-44-9 168691-45-0 168691-46-1 168691-42-7 168691-43-8 168691-49-4 168691-47-2 168691-48-3, 1-8-Gastrin-14 I (human) 168691-50-7 168691-51-8 168691-52-9 168691-53-0 168691-54-1 168691-57-4 168691-58-5 168691-59-6 168691-55-2 168691-56-3 168691-62-1 168691-63-2 168691-64-3 168691-60-9 168691-61-0 168691-68-7 168691-69-8 168691-65-4 168691-66-5 168691-67-6 168691-73-4 168691-74-5 168691-70-1 168691-71-2 168691-72-3 168691-75-6 168691-76-7 168691-77-8 168691-78-9 168691-79-0 168691-84-7 168691-80-3 168691-81-4 168691-82-5 168691-83-6 168691-85-8 168691-86-9 168691-87-0 168691-88-1 168691-89-2 168691-94-9 168691-90-5 168691-91-6 168691-92-7 168691-93-8 168691-99-4 168691-95-0 168691-96-1 168691-97-2 168691-98-3 168692-03-3 168692-04-4 168692-00-0 168692-01-1 168692-02-2 168692-05-5 168692-06-6 168692-07-7 168692-08-8 168692-09-9 168692-13-5 168692-14-6 168692-10-2 168692-11-3 168692-12-4 168692-17-9 168692-18-0 168692-19-1 168692-15-7 168692-16-8 168692-22-6 168692-23-7 168692-24-8 168692-20-4 168692-21-5 168692-27-1 168692-28-2 168692-29-3 168692-25-9 168692-26-0 168692-30-6 168692-31-7 168692-32-8 168692-33-9 168692-34-0 168692-37-3 168692-38-4 168692-39-5 168692-35-1 168692-36-2 168692-40-8 RL: BOC (Biological occurrence); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); USES (Uses) (hydrophilic signal oligopeptides and methods of therapeutic use) IT 168692-42-0 168692-43-1 168692-44-2 168692-45-3 168692-41-9 168692-48-6 168692-49-7 168692-50-0 168692-46-4 168692-47-5 168692-52-2 168692-53-3 168692-54-4 168692-55-5 168692-51-1 168692-57-7 168692-58-8 168692-59-9 168692-60-2 168692-56-6 168692-61-3 168692-62-4 168692-63-5 168692-64-6 168692-65-7 168692-66-8 168692-67-9 168692-68-0 168692-69-1 168692-70-4 168692-71-5 168692-72-6 168692-73-7 168692-74-8 168692-75-9

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168692-77-1
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        (hydrophilic signal oligopeptides and methods of therapeutic use)
    80965-96-4, Elastase, prepro-
    RL: BPR (Biological process); BSU (Biological study, unclassified); PRP
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        (of Schistosoma, signal fragments; hydrophilic signal oligopeptides and
       methods of therapeutic use)
    9028-35-7, Hydroxymethylglutaryl coenzyme A reductase 39364-01-7,
    Prorenin 50812-36-7, Synthetase, farnesyl pyrophosphate 75432-63-2,
    Glucagon, prepro- 81690-22-4, Preprogastrin 106602-62-4, Islet amyloid
    polypeptide 113834-12-1, Schistosomin
                                             123774-88-9,
    Gonadotropin-releasing factor, pro- 140208-23-7, Plasminogen
    activator inhibitor 1 142243-03-6, Plasminogen activator
    inhibitor 2
    RL: BPR (Biological process); BSU (Biological study, unclassified); PRP
     (Properties); THU (Therapeutic use); BIOL (Biological study); PROC
     (Process); USES (Uses)
        (signal fragments; hydrophilic signal oligopeptides and methods of
       therapeutic use)
    ANSWER 12 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
    1992:77826 HCAPLUS
    116:77826
    Entered STN: 06 Mar 1992
    Manufacture of fusion protein containing the kringle region of
    plasminogen
    Yokoo, Yoshiharu; Sugimoto, Seiji; Sato, Moriyuki; Nishi, Tatsuya; Ito,
    Kyowa Hakko Kogyo Co., Ltd., Japan
    Jpn. Kokai Tokkyo Koho, 18 pp.
    CODEN: JKXXAF
    Patent
    Japanese
    ICM C12P021-02
    ICS C07K003-10; C07K003-20; C07K007-10; C07K013-00
    C12N015-62; C12P021-06
    C12P021-02, C12R001-19; C07K099-00
    3-4 (Biochemical Genetics)
FAN.CNT 1
    PATENT NO.
                       KIND
                              DATE
                                        APPLICATION NO.
                                                               DATE
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                        _ _ _ _
                              -----
                                          -----
                                                                -----
    JP 03219892
                        A2
                              19910927
                                       JP 1990-13941
                                                               19900124 <--
PRAI JP 1990-13941
                              19900124 <--
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CLASS
PATENT NO.
                CLASS PATENT FAMILY CLASSIFICATION CODES
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JP 03219892
               ICM
                       C12P021-02
                ICS
                       C07K003-10; C07K003-20; C07K007-10; C07K013-00
               ·ICA
                       C12N015-62; C12P021-06
                       C12P021-02, C12R001-19; C07K099-00
                ICI
    Manufacture of the kringle region 1 (k1) of plasminogen and/or a
AB
    heterologous protein via the expression of a chimeric gene thereof is
     described. K1 and the adjacent protein, e.g. the B domain of protein A
     domain (I) of Staphylococcus aureus, are linked with an amino acid/peptide
     linker that can be easily cleaved by a chemical/enzymic treatment for
separation
    and purification A histidine residue can also be introduced into the protein
     as chelating sites. Plasmids pPrKT1 and pPZKT1 encoding the fusion
    protein of k1-I and k1-I derivative, resp., for expression in Escherichia coli
    were given. Purifn, of the fusion protein with lysine-affinity chromatog.
    was also shown.
    kringle 1 plasminogen fusion protein; protein A
    plasminogen kringle 1
IT
    Animal cell
    Escherichia coli
        (expression in, of chimeric gene for human plasminogen
       kringle 1 and Staphylococcus aureus)
IT
    Microorganism
        (expression in, of chimeric gene for human plasminogen
       kringle 1 and Staphylococcus aureus protein A B domain)
TΤ
     Plasmid and Episome
        (pPZKT1, chimeric gene for human plasminogen kringle 1 and
        Staphylococcus aureus protein A B domain on,)
IT
     Plasmid and Episome
        (pPrKT1, chimeric gene for human plasminogen kringle 1 and
        Staphylococcus aureus protein A B domain on,)
     Proteins, specific or class
IT
     RL: BIOL (Biological study)
        (A, B domain, fusion products with kringle 1 of plasminogen
       of, recombinant manufacture of)
     91931-07-6, 212-269-Protein A (Staphylococcus aureus clone pAC37)
IT
     138726-05-3, 80-165-Plasminogen (human liver clone
     pPLGKG protein moiety reduced)
    RL: PRP (Properties)
        (amino acid sequence of, fusion protein containing)
     71-00-1, Histidine, biological studies
IT
    RL: BIOL (Biological study)
        (chelating site, in fusion protein containing plaminogen kringle 1 and B
       domain of protein A)
IT
     56-87-1, Lysine, analysis
     RL: ANST (Analytical study)
        (kringle 1 carboxyl terminus of plasminogen containing, purification
       by affinity chromatog. of)
IT
     9001-91-6P, Plasminogen
    RL: PREP (Preparation)
        (kringle 1 domain of, fusion products with B domain of protein A and,
        recombinant preparation of)
L36
    ANSWER 13 OF 13 HCAPLUS COPYRIGHT 2005 ACS on STN
     1989:491713 HCAPLUS
AN
DN
     111:91713
    Entered STN: 16 Sep 1989
ED
    Manufacture of tissue plasminogen activator analogs by
TI
    recombinant DNA technology
    Mulvihill, Eileen R.; Nexo, Bjorn A.; Yoshitake, Shinji; Ikeda, Yasunori;
```

Suzuki, Suguru; Hashimoto, Akira; Yuzuriha, Teruaki

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Zymogenetics, Inc., USA; Novo Industri A/S; Eisai Co., Ltd.
PΑ
SO
     Eur. Pat. Appl., 54 pp.
     CODEN: EPXXDW
DT
     Patent
     English
LA
IC
     ICM C12N009-50
     ICS C12N015-00; C07H021-04; C12N005-00; A61K037-54
ICI
     C12N005-00, C12R001-19
     3-4 (Biochemical Genetics)
     Section cross-reference(s): 13, 63
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                                          APPLICATION NO. DATE
     PATENT NO.
                        KIND DATE
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                                              _____
                                                                      EP 293934
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                          A1
                                 19881207 EP 1988-108949 19880603 <--
     EP 293934
                          B1
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         R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE
     R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE
FI 8802628

NO 8802453

A 19881205

NO 179754

B 19960902

NO 179754

C 19961211

DK 8803022

A 19890203

DK 1988-3022

ZA 8803958

A 19890726

ES 2058180

T3 19941101

ES 1988-108949

JP 01085078

A2 19890330

JP 1988-138232

JP 04048433

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KR 9705251

B1 19970414

KR 1988-6716

AU 8817410

AU 617323

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     AU 617323
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                                 19920922
                                            US 1991-747452
                                                                     19910812 <--
PRAI US 1987-58217
                         Α
                                 19870604 <--
 PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES
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 EP 293934
                ICM
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                 ICS
                         C12N015-00; C07H021-04; C12N005-00; A61K037-54
                 ICI
                         C12N005-00, C12R001-19
 EP 293934
                 ECLA
                         C12N009/72B
                                                                                <--
     Tissue plasminogen activator (t-PA) analogs which exhibit
     greater specificity for fibrin than native t-PA are disclosed. Native
     t-PA contains 2 triple disulfide-bonded regions, the kringle domains K1
     and K2, which participate in the binding of t-PA to fibrin. In this
     invention, the K1 domain of native t-PA is replaced with that from another
     source. The t-PA analogs may further include a variety of substitutions
     and modifications. A cDNA comprising the coding sequence for native human
     t-PA was constructed from an mRNA isolated from the Bowes melanoma cell
     line. This cDNA was then used to construct the plasmid pDR1296. Because
     the prepro-sequence of t-PA was not present in pDR1296, it was constructed
     from synthesized oligonucleotides and subsequently joined to the cDNA in
     the vector Zem99. Here the complete t-PA coding sequence was sandwiched
     between a metallothionein I promoter and a human growth hormone
     terminator. The K1 domain of plasminogen was constructed from
     11 oligonucleotides and was then inserted into the t-PA cDNA as a
     replacement for the K1 domain of t-PA. The resultant plasmid, Zem99-8000,
     was used to transform E. coli.
ST
     tissue plasminogen activator mutation; human tissue
     plasminogen activator gene cloning
IT
     Mammal
        (cloning in cells of, of tissue plasminogen activator analog
        gene of human)
IT
     Escherichia coli
        (cloning in, of tissue plasminogen activator analog gene of
        human)
```

IT

Gene and Genetic element, animal

```
RL: BIOL (Biological study)
        (for tissue plasminogen activator analog, of human)
IT
     Protein sequences
        (of kringle domain for human tissue plasminogen activator
        analog)
IT
     Molecular cloning
        (of tissue plasminogen activator analog gene, of human)
IT
     Protein sequences
        (of tissue plasminogen activator native and variants forms,
        of human, complete)
ΙT
     Fibrins
     RL: BIOL (Biological study)
        (tissue plasminogen activator of human with enhanced binding
        to, construction of)
ΙT
     Proteins, specific or class
     RL: PRP (Properties)
        (C, growth factor domain of, in tissue plasminogen activator
        analog of human)
IT
     Plasmid and Episome
        (Zem99, tissue plasminogen activator gene of human on, for
        site-specific mutagenesis)
     Plasmid and Episome
IT
        (Zem99-8000, tissue plasminogen activator analog gene of
        human on)
     Plasmid and Episome
TT
        (Zem99-8100, tissue plasminogen activator analog gene of
        human on)
IT
     Plasmid and Episome
        (pDR1296, tissue plasminogen activator gene of human on,
        cloning of, in Escherichia coli)
     Deoxyribonucleic acid sequences
IT
        (plasmin-specifying, kringle domain)
IT
     Mutation
        (site-specific, of tissue plasminogen activator, of human,
        for enhanced binding to fibrin)
     Deoxyribonucleic acid sequences
IT
        (tissue-type plasminogen activator-specifying, native and
        variant forms, of human, complete)
IT
     122007-78-7
                   122007-79-8
                                 122007-80-1
                                                122007-81-2
                                                              122007-82-3
     122007-83-4
                   122007-84-5
                                 122007-85-6
     RL: PRP (Properties)
        (amino acid sequence of)
IT
     84933-03-9, Plasminogen activator (human tissue-type precursor
     protein moiety reduced)
                               84933-04-0, Plasminogen activator
     (human tissue-type protein moiety reduced)
     RL: PRP (Properties)
        (amino acid sequence of, preparation of analogs of)
ΙT
     122071-87-8, 84-162-Plasminogen (human liver clone
     pPLGKG protein moiety reduced)
     RL: PRP (Properties)
        (amino acid sequence of, recombinant tissue plasminogen
        activator containing)
                                  122071-60-7
                                                              122071-62-9
ΙT
     122071-58-3
                   122071-59-4
                                                122071-61-8
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                                 122092-15-3
                                                122092-16-4
     122071-63-0
     RL: PRP (Properties)
        (as finger domain of human tissue plasminogen activator
        analog)
     122071-86-7
IT
     RL: PRP (Properties)
        (as kringle domain for human tissue plasminogen activator
        analog)
     9001-25-6, Blood-coagulation factor VII 9001-28-9, Factor IX
IT
     9001-29-0, Factor X
```

RL: PRP (Properties) (growth factor domain of, in tissue plasminogen activator analog of human) 9001-26-7, Prothrombin 9001-30-3, Blood-coagulation factor XII TΤ 9001-91-6, Plasminogen RL: PRP (Properties) (kringle domain of, substitution of, for that of human tissue plasminogen activator) 122006-82-0, Deoxyribonucleic acid (human clone Zem94 TT 122006-81-9 tissue-type plasminogen activator messenger RNA-complementary) 122006-87-5 122006-86-4 122006-88-6 122006-89-7 122006-83-1 RL: PRP (Properties); BIOL (Biological study) (nucleotide sequence of) 105913-11-9, Plasminogen activator IT RL: PRP (Properties) (tissue-type, of human, replacement of kringle domain of, for enhanced binding to fibrin) => fil reg FILE 'REGISTRY' ENTERED AT 14:01:47 ON 15 FEB 2005 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2005 American Chemical Society (ACS) Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem. STRUCTURE FILE UPDATES: 14 FEB 2005 HIGHEST RN 831169-46-1 DICTIONARY FILE UPDATES: 14 FEB 2005 HIGHEST RN 831169-46-1 TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004 Please note that search-term pricing does apply when conducting SmartSELECT searches. Crossover limits have been increased. See HELP CROSSOVER for details. Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html => => d browse 138 :1-51 L38 ANSWER 1 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN RN 679784-39-5 REGISTRY L-Aspartic acid, L-lysyl-L-leucyl-L-tyrosyl- (9CI) (CA INDEX NAME) CN FS PROTEIN SEQUENCE; STEREOSEARCH C25 H39 N5 O8 MF SR CA LC STN Files: CA, CAPLUS DT.CA CAplus document type: Journal Roles from non-patents: BIOL (Biological study); PRP (Properties); USES RL.NP (Uses) **RELATED SEQUENCES AVAILABLE WITH SEQLINK**

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 2 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 679784-35-1 REGISTRY

CN L-Aspartic acid, N2-acetyl-L-lysyl-L-leucyl-L-tyrosyl- (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C27 H41 N5 O9

SR CA

LC STN Files: CA, CAPLUS

DT.CA CAplus document type: Journal

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 1 REFERENCES IN FILE CA (1907 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L38 ANSWER 3 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
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- CN 38: PN: US6699838 SEQID: 38 unclaimed protein (9CI) (CA INDEX NAME)
- FS PROTEIN SEQUENCE
- MF Unspecified
- CI MAN
- SR CA
- LC STN Files: CA, CAPLUS, USPATFULL

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       Roles from patents: PRP (Properties)
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L38 ANSWER 4 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
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CN
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MF
     Unspecified
CI
     MAN
SR
     CA
LC
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                  CA, CAPLUS, USPATFULL
DT.CA CAplus document type: Patent
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                  CA, CAPLUS, USPATFULL
LC
DT.CA CAplus document type: Patent
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               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
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L38
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DT.CA CAplus document type: Patent
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               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
L38 ANSWER 8 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
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       Roles from patents: BIOL (Biological study); PREP (Preparation); PRP
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Absolute stereochemistry.
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PAGE 1-B

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 10 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 506450-18-6 REGISTRY

CN Plasminogen (cattle kringle 1 fragment) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 13: PN: US20030064926 SEQID: 11 claimed protein

FS PROTEIN SEQUENCE

MF Unspecified

CI MAN

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent.

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

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L38 ANSWER 11 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 506450-15-3 REGISTRY

CN Plasminogen (human kringle 1 fragment) (9CI) (CA INDEX NAME)

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RL.P
       (Properties)
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PAGE 1-B

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L38 ANSWER 14 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 265296-51-3 REGISTRY

CN 33: PN: US6057122 SEQID: 38 unclaimed protein (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE

MF Unspecified

CI MAN

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: PRP (Properties)

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L38 ANSWER 15 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 265296-50-2 REGISTRY

CN 32: PN: US6057122 SEQID: 37 unclaimed protein (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE

MF Unspecified

CI MAN

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: PRP (Properties)

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1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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L38 ANSWER 17 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
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                 CA, CAPLUS, TOXCENTER, USPATFULL
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DT.CA CAplus document type: Patent
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     265110-89-2 REGISTRY
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     C45 H67 N11 O12
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SR
                 CA, CAPLUS, TOXCENTER, USPATFULL
LC
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DT.CA CAplus document type: Patent
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PAGE 1-B

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1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 19 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 264868-26-0 REGISTRY

CN 355-543-Plasminogen (human) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 7: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE

MF Unspecified

CI MAN

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PROC (Process); PRP (Properties)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

*** USE 'SQD' OR 'SQIDE' FORMATS TO DISPLAY SEQUENCE ***

1 REFERENCES IN FILE CA (1907 TO DATE)

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1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 20 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 251555-93-8 REGISTRY

CN L-Aspartic acid, N2-[(1,1-dimethylethoxy)carbonyl]-L-lysyl-L-leucyl-N-methyl-L-phenylalanyl-, bis(phenylmethyl) ester (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C45 H61 N5 O9

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER DT.CA CAplus document type: Patent

RL.P Roles from patents: PREP (Preparation); RACT (Reactant or reagent)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 21 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 251555-92-7 REGISTRY

CN L-α-Asparagine, N6-[(phenylmethoxy)carbonyl]-L-lysyl-L-leucyl-Nmethyl-O-(phenylmethyl)-L-tyrosyl-, 1,1-dimethylethyl ester (9CI) (CA
INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C45 H62 N6 O9

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER

DT.CA CAplus document type: Patent

RL.P Roles from patents: PREP (Preparation); RACT (Reactant or reagent)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

- 1 REFERENCES IN FILE CA (1907 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 22 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 251555-91-6 REGISTRY

CN $L-\alpha$ -Asparagine, N2-[(1,1-dimethylethoxy)carbonyl]-N6-[(phenylmethoxy)carbonyl]-L-lysyl-L-leucyl-N-methyl-O-(phenylmethyl)-L-tyrosyl-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C50 H70 N6 O11

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER

DT.CA CAplus document type: Patent

RL.P Roles from patents: PREP (Preparation); RACT (Reactant or reagent)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 23 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 251555-86-9 REGISTRY

CN L-Aspartic acid, N2-[(1,1-dimethylethoxy)carbonyl]-L-lysyl-N-methyl-L-leucyl-ψ(CH2-NH)-L-phenylalanyl- (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C31 H51 N5 O8

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER

DT.CA CAplus document type: Patent

RELATED SEQUENCES AVAILABLE WITH SEQLINK

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 S
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 Me
 NH_2
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 OBu

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 24 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 251555-85-8 REGISTRY

CN L-Aspartic acid, N2-[(1,1-dimethylethoxy)carbonyl]-L-lysyl-L-leucyl-N-methyl-L-tyrosyl-N-methyl- (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C32 H51 N5 O10

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LC STN Files: CA, CAPLUS, TOXCENTER

DT.CA CAplus document type: Patent

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RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

$$CO_2H$$
 CO_2H
 C

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1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 25 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN **251555-84-7** REGISTRY

CN L-Aspartic acid, N2-[(1,1-dimethylethoxy)carbonyl]-L-lysyl-N-methyl-L-leucyl-N-methyl-L-tyrosyl- (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C32 H51 N5 O10

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RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

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1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 26 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 251555-83-6 REGISTRY

CN L-Aspartic acid, N2-acetyl-L-lysyl-L-leucyl-N-methyl-L-phenylalanyl- (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C28 H43 N5 O8

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LC STN Files: CA, CAPLUS, TOXCENTER

DT.CA CAplus document type: Patent

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- L38 ANSWER 27 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
- RN 251555-82-5 REGISTRY
- CN L-α-Asparagine, N2-acetyl-L-lysyl-L-leucyl-N-methyl-L-tyrosyl- (9CI)

(CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C28 H44 N6 O8

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LC STN Files: CA, CAPLUS, TOXCENTER

DT.CA CAplus document type: Patent

RELATED SEQUENCES AVAILABLE WITH SEQLINK

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L38 ANSWER 28 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 250163-86-1 REGISTRY

CN 11: PN: US5981484 SEQID: 11 unclaimed protein (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE

MF Unspecified

CI MAN

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: PRP (Properties)

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1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 29 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 250159-84-3 REGISTRY

CN 454-546-Plasminogen (human) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 6: PN: US6057122 TABLE: 1 claimed protein

CN 7: PN: US5981484 SEQID: 7 claimed protein

FS PROTEIN SEQUENCE

MF Unspecified

CI MAN

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LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

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L38 ANSWER 35 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
     199664-91-0 REGISTRY
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     L-tyrosyl-L-\alpha-aspartyl- (9CI) (CA INDEX NAME)
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     PROTEIN SEQUENCE; STEREOSEARCH
MF
     C47 H69 I N12 O12
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     STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
DT.CA CAplus document type: Patent
       Roles from patents: BIOL (Biological study); PREP (Preparation); PRP
       (Properties); USES (Uses)
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RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

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3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 36 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-90-9 REGISTRY

CN L-Tyrosinamide, 1-acetyl-L-prolyl-L-arginyl-L-lysyl-L-leucyl-L-tyrosyl-L- α -aspartyl-3-(iodo-1251)- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 20: PN: US6057122 PAGE: 39/40 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C47 H69 I N12 O12

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

PAGE 1-B

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3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 37 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-89-6 REGISTRY

CN L- α -Asparagine, N2-acetyl-L-lysyl-L-leucyl-L-tyrosyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 19: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C27 H42 N6 O8

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 38 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-88-5 REGISTRY

CN L-Tyrosinamide, 1-acetyl-L-prolyl-L-arginyl-L-lysyl-L-leucyl-L-tyrosyl-L- $\alpha\text{-aspartyl-3-iodo-}$ (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 18: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C47 H69 I N12 O12

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

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3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 39 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-87-4 REGISTRY

CN L-Tyrosinamide, 1-acetyl-L-prolyl-L-arginyl-L-lysyl-L-leucyl-3-iodo-L-tyrosyl-L- α -aspartyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 17: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C47 H69 I N12 O12

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

PAGE 1-B

3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 40 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-86-3 REGISTRY

CN L-Tyrosinamide, N2-acetyl-L-arginyl-L-lysyl-L-leucyl-L-tyrosyl-L- α -aspartyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 16: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C42 H63 N11 O11

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 41 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-85-2 REGISTRY

CN L-Tyrosinamide, 1-acetyl-L-prolyl-L-α-glutamyl-L-lysyl-L-arginyl-L-

tyrosyl-L-α-aspartyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 15: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C46 H66 N12 O14

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

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3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 42 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-84-1 REGISTRY

CN L- α -Asparagine, 1-acetyl-L-prolyl-L-arginyl-L-lysyl-L-leucyl-L-tyrosyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 14: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C38 H61 N11 010

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 43 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 199664-83-0 REGISTRY

CN L-Tyrosinamide, 1-acetyl-L-prolyl-L-arginyl-L-lysyl-L-leucyl-L-tyrosyl-L- α -aspartyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN 13: PN: US6057122 TABLE: 1 claimed protein

FS PROTEIN SEQUENCE; STEREOSEARCH

MF C47 H70 N12 O12

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation); PRP (Properties)

RELATED SEQUENCES AVAILABLE WITH SEQLINK

Absolute stereochemistry.

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- 4 REFERENCES IN FILE CA (1907 TO DATE)
- 4 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L38 ANSWER 44 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
- RN 199664-82-9 REGISTRY
- CN L-Tyrosinamide, N-acetyl-L-tyrosyl-L-threonyl-L-threonyl-L-asparaginyl-Lprolyl-L-arginyl-L-lysyl-L-leucyl-L-tyrosyl-L-α-aspartyl- (9CI) (CA
 INDEX NAME)

OTHER NAMES:

- CN 12: PN: US6057122 TABLE: 1 claimed protein
- FS PROTEIN SEQUENCE; STEREOSEARCH
- MF C68 H99 N17 O20
- SR CA
- LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
- DT.CA CAplus document type: Journal; Patent
- RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)
- RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation); PRP (Properties)
- **RELATED SEQUENCES AVAILABLE WITH SEQLINK**

Absolute stereochemistry.

PAGE 1-B

$$(CH_2)_3 \xrightarrow{H} NH_2$$

$$S \xrightarrow{H} NH$$

$$S \xrightarrow{H}$$

PAGE 1-C

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4 REFERENCES IN FILE CA (1907 TO DATE)

4 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 45 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 196417-08-0 REGISTRY

CN Plasminogen (human kringle 5 domain-containing fragment) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Plasminogen (human blood kringle 5 domain 80-amino-acid fragment)

FS PROTEIN SEQUENCE

MF Unspecified

CI MAN

SR CA

LC STN Files: CA, CAPLUS

DT.CA CAplus document type: Journal

RL.NP Roles from non-patents: BIOL (Biological study); PROC (Process); PRP (Properties)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

*** USE 'SQD' OR 'SQIDE' FORMATS TO DISPLAY SEQUENCE ***

2 REFERENCES IN FILE CA (1907 TO DATE)

2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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L38 ANSWER 46 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
     185074-41-3 REGISTRY
RN
CN
     Angiostatin (cattle krinkle 1 region-contg. fragment) (9CI) (CA INDEX
     NAME)
OTHER CA INDEX NAMES:
     Angiostatin (ox krinkle 1 region-contg. fragment)
CN
     PROTEIN SEQUENCE
FS
MF
     Unspecified
CI
     MAN
SR
     CA
                  CA, CAPLUS, TOXCENTER
LC
     STN Files:
DT.CA CAplus document type: Patent
       Roles from patents: BIOL (Biological study); PRP (Properties); USES
RL.P
       (Uses)
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               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
    ANSWER 47 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
L38
     185074-38-8 REGISTRY
RN
     Angiostatin (mouse krinkle 1 region-contg. fragment) (9CI) (CA INDEX
CN
     NAME)
     PROTEIN SEQUENCE
FS
     Unspecified
MF
CI
     MAN
SR
                  CA, CAPLUS, TOXCENTER
LC
     STN Files:
DT.CA CAplus document type: Patent
       Roles from patents: BIOL (Biological study); PRP (Properties); USES
       (Uses)
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
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               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
L38
    ANSWER 48 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
     168693-32-1 REGISTRY
RN
     L-Histidine, L-histidyl-L-lysyl-L-leucyl-L-phenylalanyl-L-α-aspartyl-
CN
     L-alanyl-L-seryl-L-α-aspartyl-L-seryl-L-seryl-L-seryl-L-tyrosyl-L-
     lysyl- (9CI) (CA INDEX NAME)
FS
     PROTEIN SEQUENCE; STEREOSEARCH
MF
     C71 H104 N20 O24
SR
     CA
LC
     STN Files:
                  CA, CAPLUS, USPATFULL
DT.CA CAplus document type: Patent
       Roles from patents: BIOL (Biological study); OCCU (Occurrence); PRP
       (Properties); USES (Uses)
Absolute stereochemistry.
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PAGE 1-C

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- 1 REFERENCES IN FILE CA (1907 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L38 ANSWER 49 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN

RN 138726-05-3 REGISTRY

CN 80-165-Plasminogen (human liver clone pPLGKG protein moiety reduced) (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE

MF C423 H635 N121 O144 S7

CI MAN

SR CA

LC STN Files: CA, CAPLUS

DT.CA CAplus document type: Patent

RL.P Roles from patents: PRP (Properties)

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               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
    ANSWER 50 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
L38
RN
     122071-87-8 REGISTRY
CN
     84-162-Plasminogen (human liver clone pPLGKG protein moiety reduced) (9CI)
     (CA INDEX NAME)
OTHER NAMES:
     78-156-Plasminogen (human kringle 1 domain-containing fragment)
CN
     Angiostatin (human krinkle 1 region-contg. fragment)
FS
     PROTEIN SEQUENCE
MF
     C385 H582 N114 O127 S7
CI
     MAN
SR
     CA
LC
     STN Files:
                 CA, CAPLUS, TOXCENTER, USPATFULL
DT.CA CAplus document type: Journal; Patent
       Roles from patents: BIOL (Biological study); PRP (Properties); USES
RL.NP
      Roles from non-patents: BIOL (Biological study); PREP (Preparation);
       PROC (Process)
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
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               3 REFERENCES IN FILE CAPLUS (1907 TO DATE)
L38 ANSWER 51 OF 51 REGISTRY COPYRIGHT 2005 ACS on STN
     122071-86-7 REGISTRY
RN
     84-162-Plasminogen (human liver clone pPLGKG protein moiety reduced),
     88-L-aspartic acid- (9CI)
                               (CA INDEX NAME)
FS
     PROTEIN SEQUENCE
MF
    C385 H581 N113 O128 S7
CI
    MAN
SR
    CA
LC
    STN Files:
                  CA, CAPLUS, TOXCENTER, USPATFULL
DT.CA CAplus document type: Patent
      Roles from patents: PRP (Properties)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
*** USE 'SQD' OR 'SQIDE' FORMATS TO DISPLAY SEQUENCE ***
               1 REFERENCES IN FILE CA (1907 TO DATE)
               1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
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